On Epigenomic Privacy: Tracking Personal MicroRNA Expression Profiles over Time

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<u>Epigenetics</u> *"epi":* above, over (greek) *"genetics":* origin (greek)

Definition: study of cellular and phenotypic trait variations stemming from other causes than changes in the genotype Definition: small non-coding RNA molecules that regulate gene expression in plants/animals 60% of genes coding human proteins are regulated by miRNAs

external factors such as: in-utero and childhood development, environmental chemicals, aging, diet.



MicroRNA (miRNA) discovered in the early 1990s

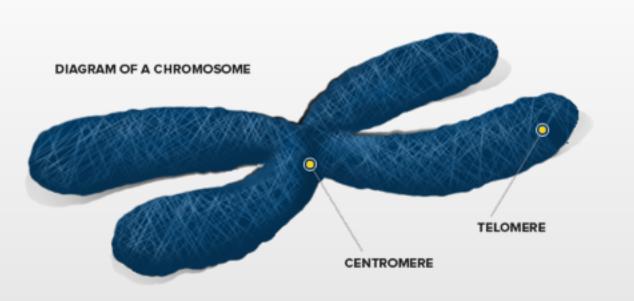
<u>MicroRNA</u> Expression Profiles

Real-valued number quantifying whether and how much miRNAs are active in a given set of cells/tissue.

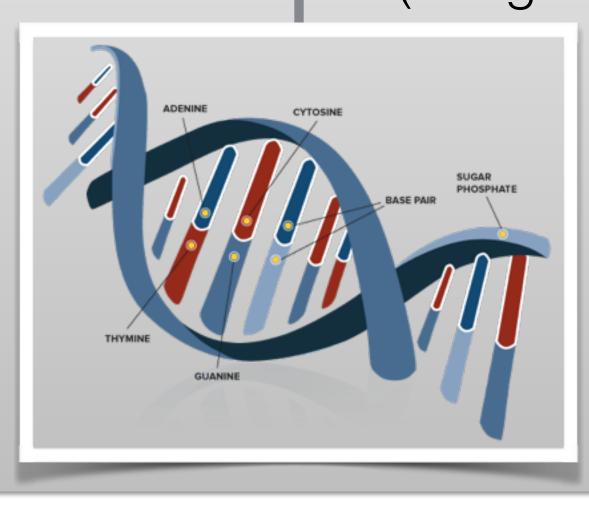




What is the purpose of MicroRNAs?



Chromosomes: carry hereditary information in long strings of **DNA** called **genes** (a region of DNA)



Graphics: genographic.nationalgeographic.com





But all cells have the same genes!



What makes the cells different: gene expression (which genes are active in a cell)



What is the purpose of MicroRNAs?

gene expression

What makes the cells different: (which genes are active in a cell)

miRNAs regulate most of human genes!

→ important for normal and **disease** cells

neurodegenerative diseases (e.g., Alzheimer's) heart diseases, diabetes, majority of cancers





RED BLOOD CELLS

More on DNA and MicroRNAs! DNA

- contains receipts what a cell potentially can do,
- is (mostly) fixed over time,
- can hint on risks of getting a disease,
- has been researched a lot. \bullet





miRNAs

- expression regulates what a cell really does,
- expression changes over time,
- can tell whether you carry a disease,
- so far, have been largely overlooked (in privacy)!

scope.seval(attr.change);
forEach(selectedTranscludes, function(selectedTransclude) {
 selectedScope = scope.Snew();
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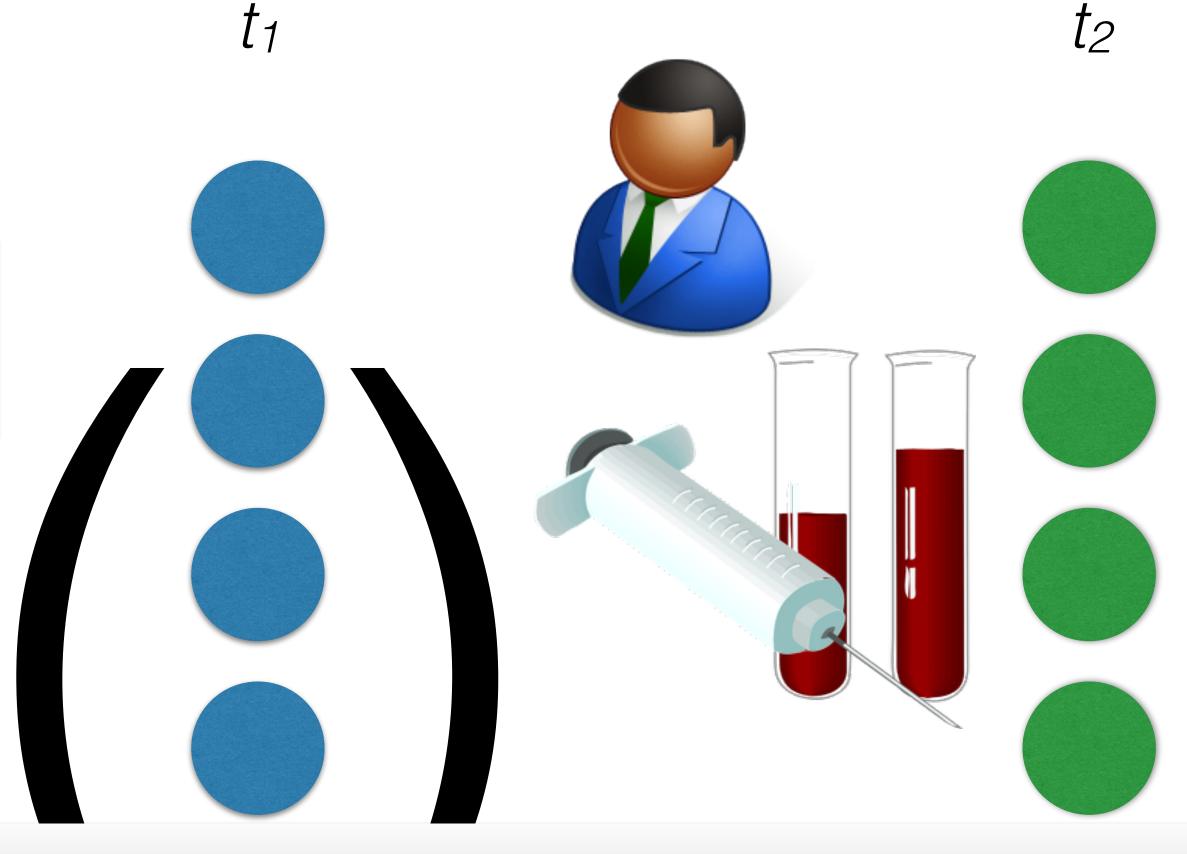


Common belief: no privacy threats from miRNAs, because of temporal variability

identification

matching

hospital server







black market

cyber attacks against healthcare companies have increased by 72% within one year





Athletes' dataset



Participants: 29 Time shift: 1 week Disease: **none**

blood-based

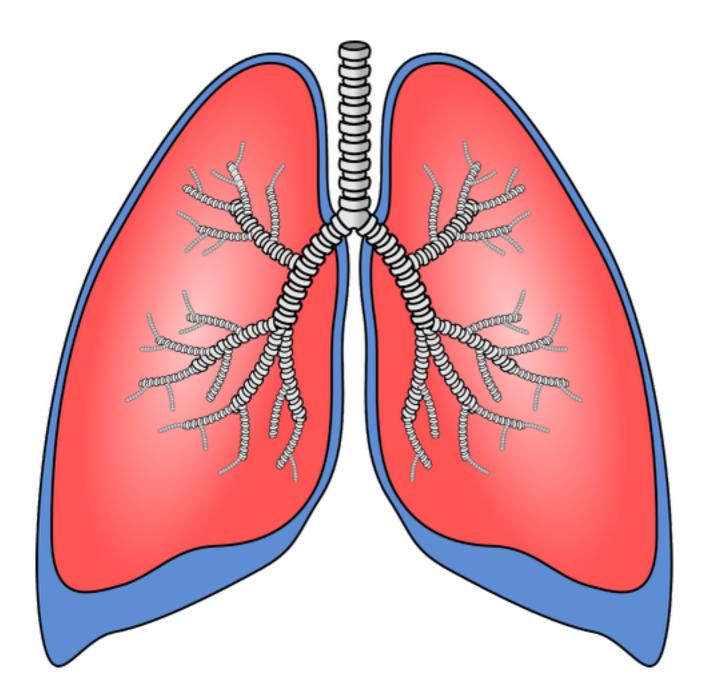


Points in time: 2 (before and after exercising)

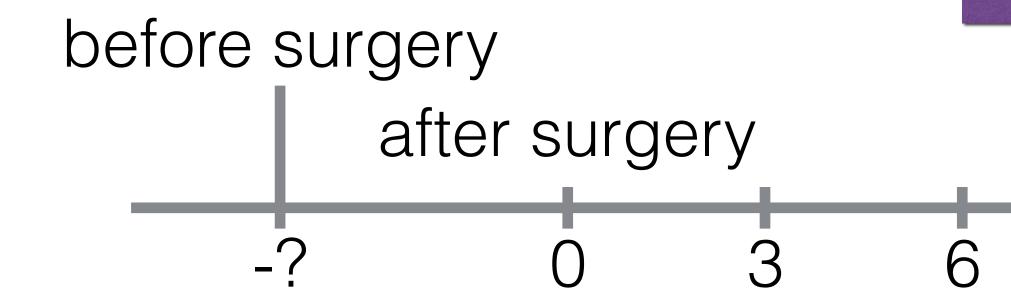
plasma-based

1,189 miRNAs per sample

Lung cancer dataset



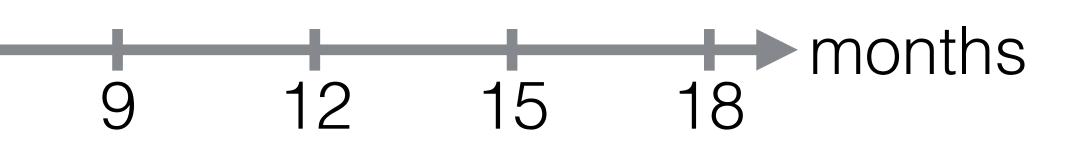
Participants: **26** Points in time: **8** Time shift: mostly **3 months** Disease: **lung cancer**



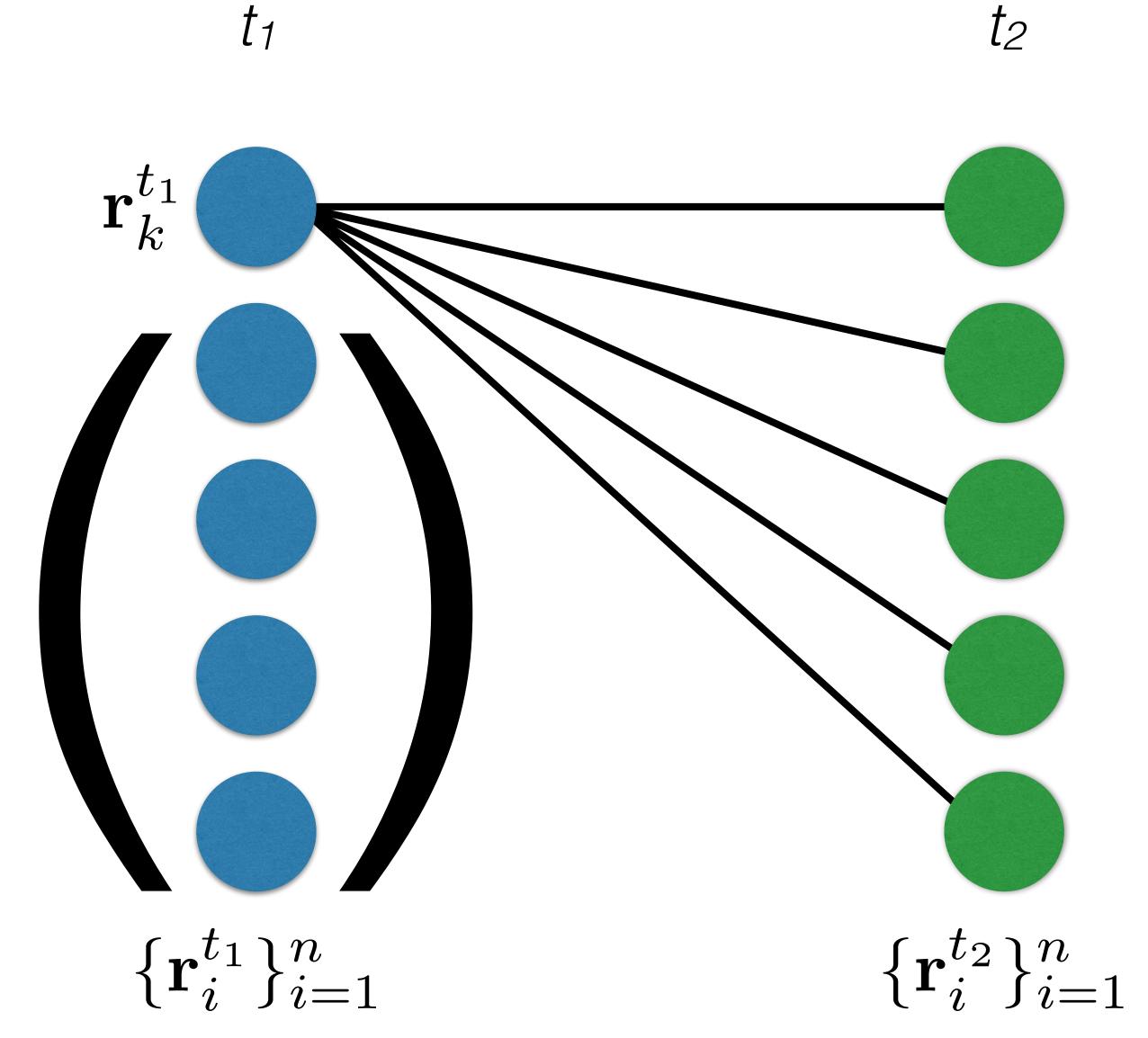


plasma-based

1,189 miRNAs per sample



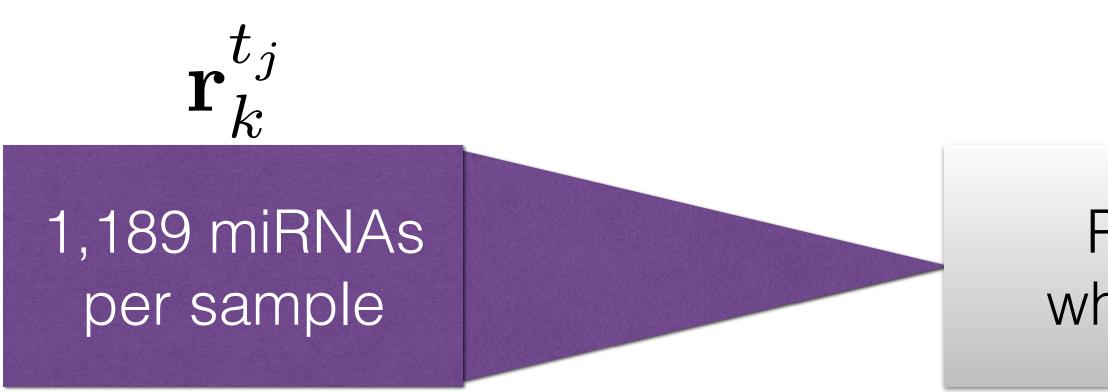






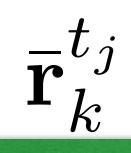
1,189 miRNAs per sample





whitening: unit variance PCA:





vector with m dimensions

PCA + whitening

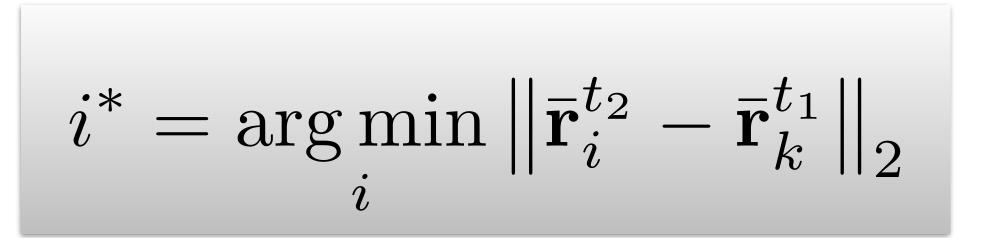
smaller dimensionality m + uncorrelated components



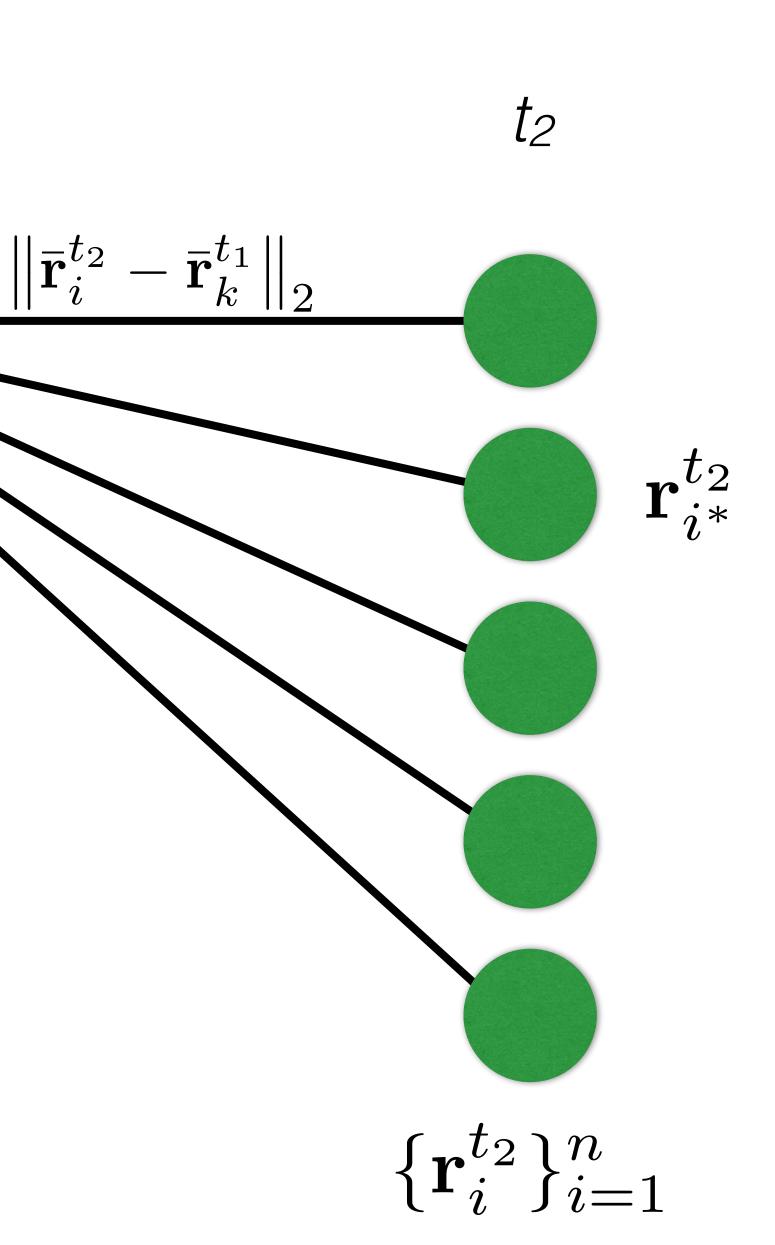


t1

 $\mathbf{r}_k^{t_1}$





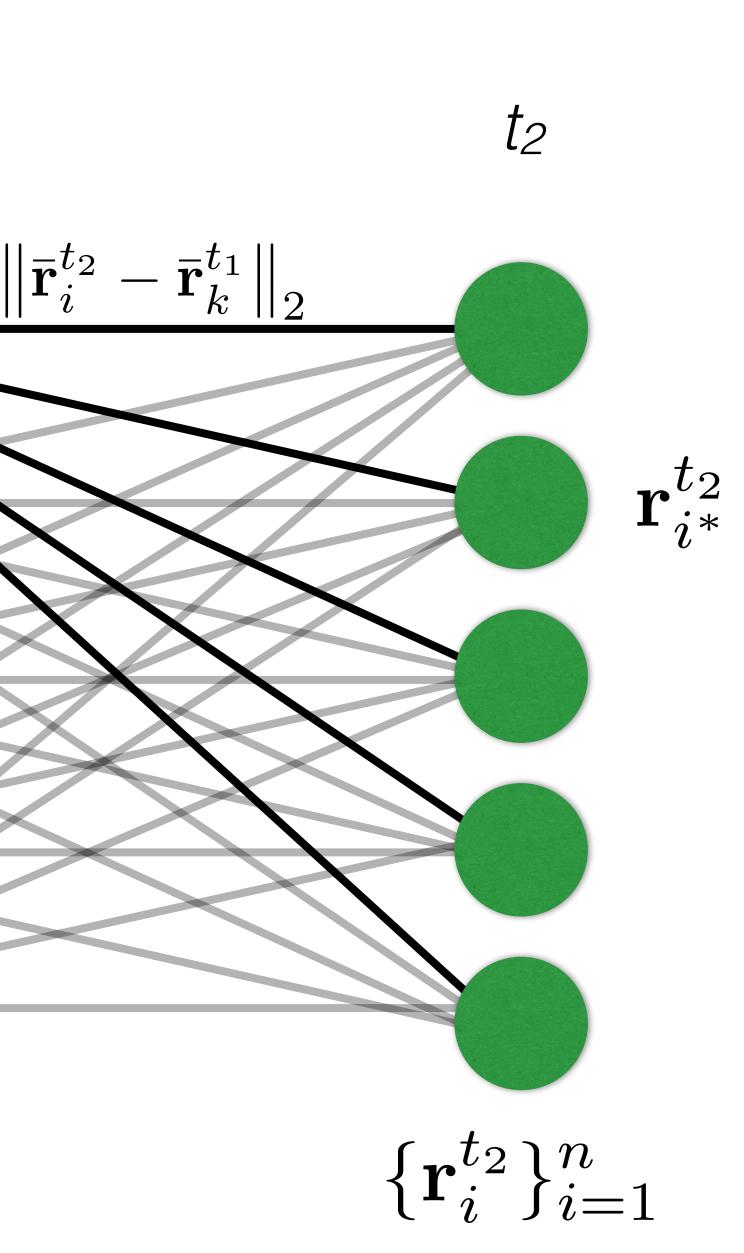


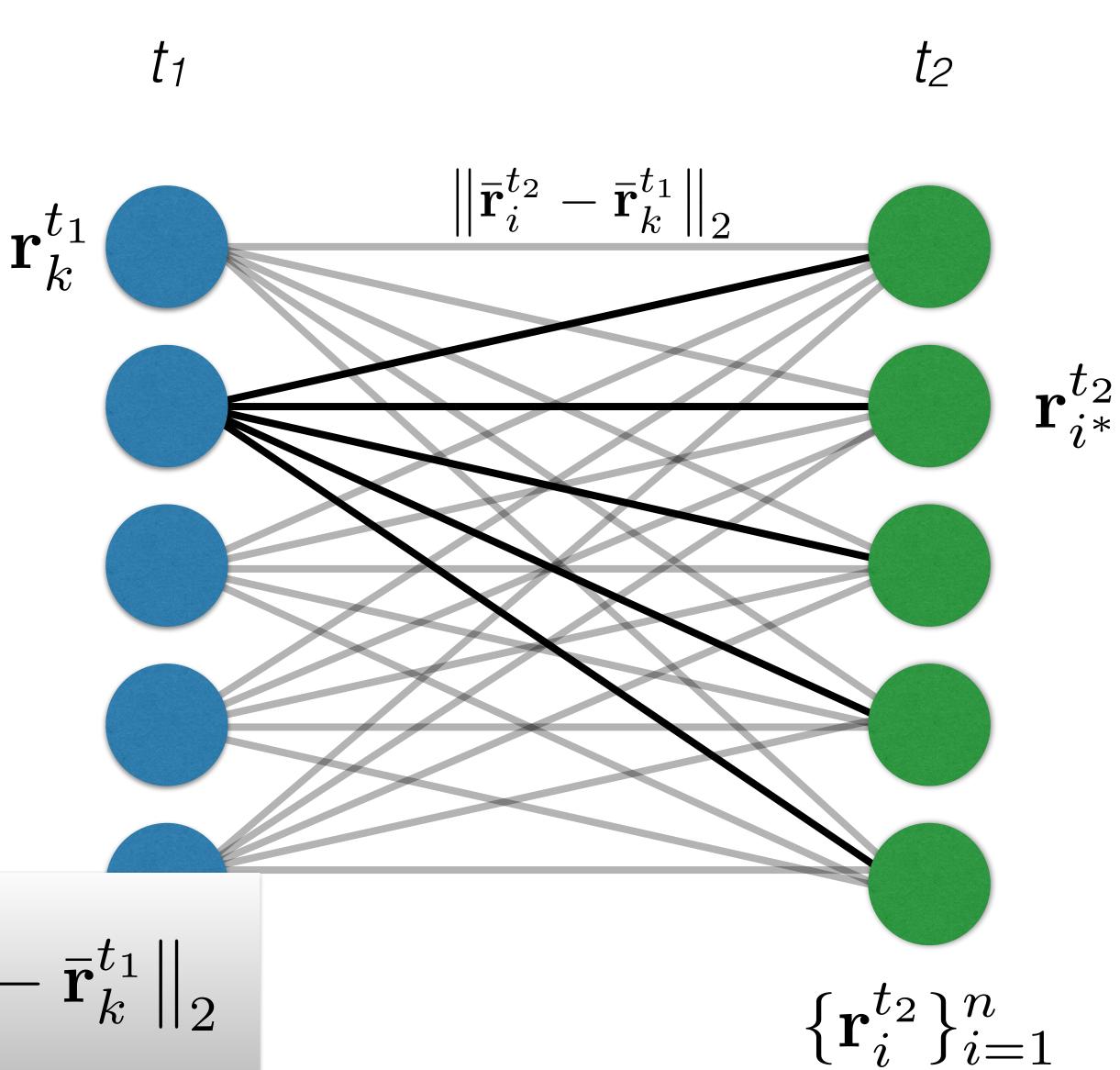
t₁

 $\mathbf{r}_k^{t_1}$

 $i^* = \underset{i}{\operatorname{arg\,min}} \|\bar{\mathbf{r}}_i^{t_2} - \bar{\mathbf{r}}_k^{t_1}\|_2$

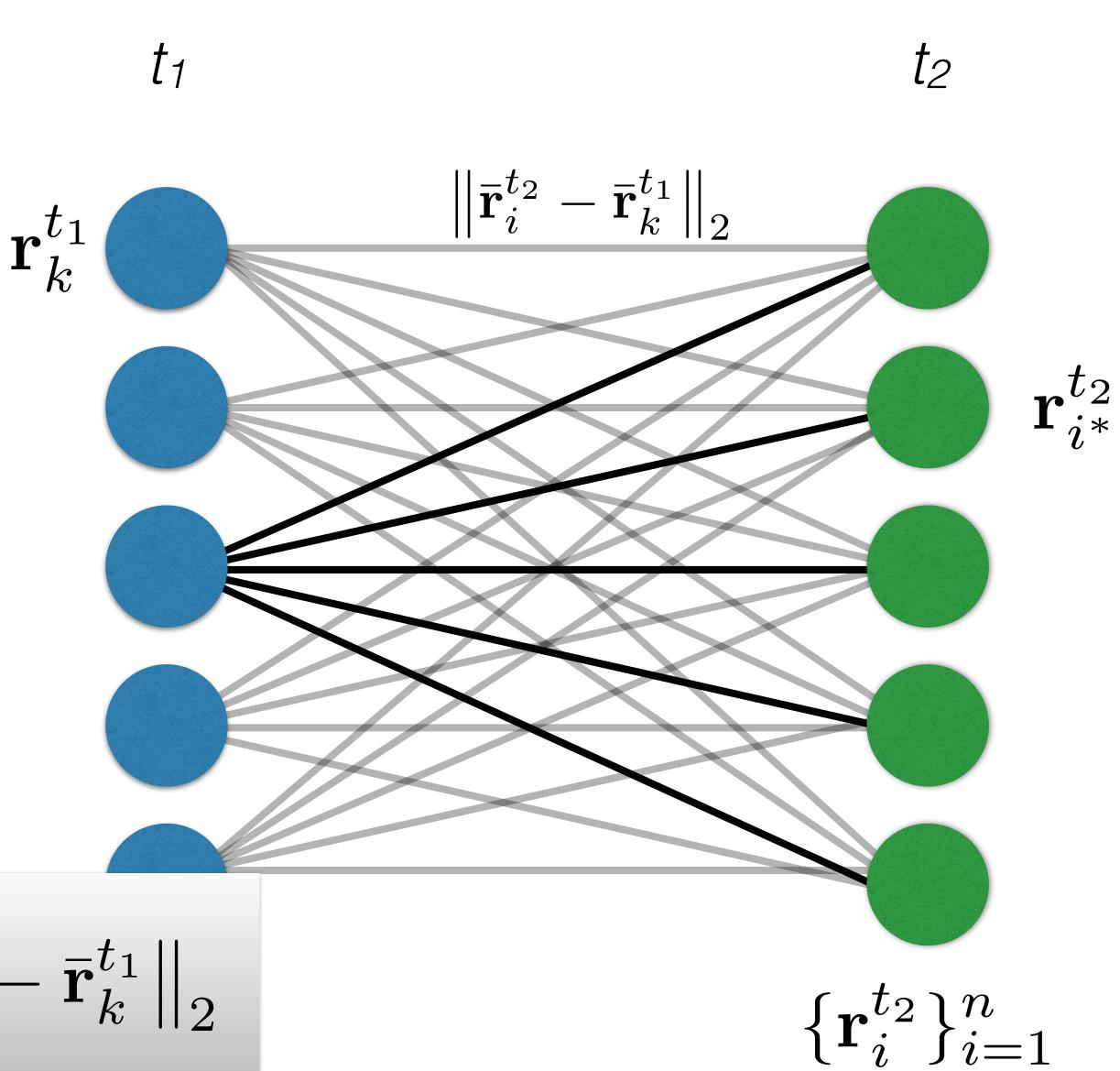






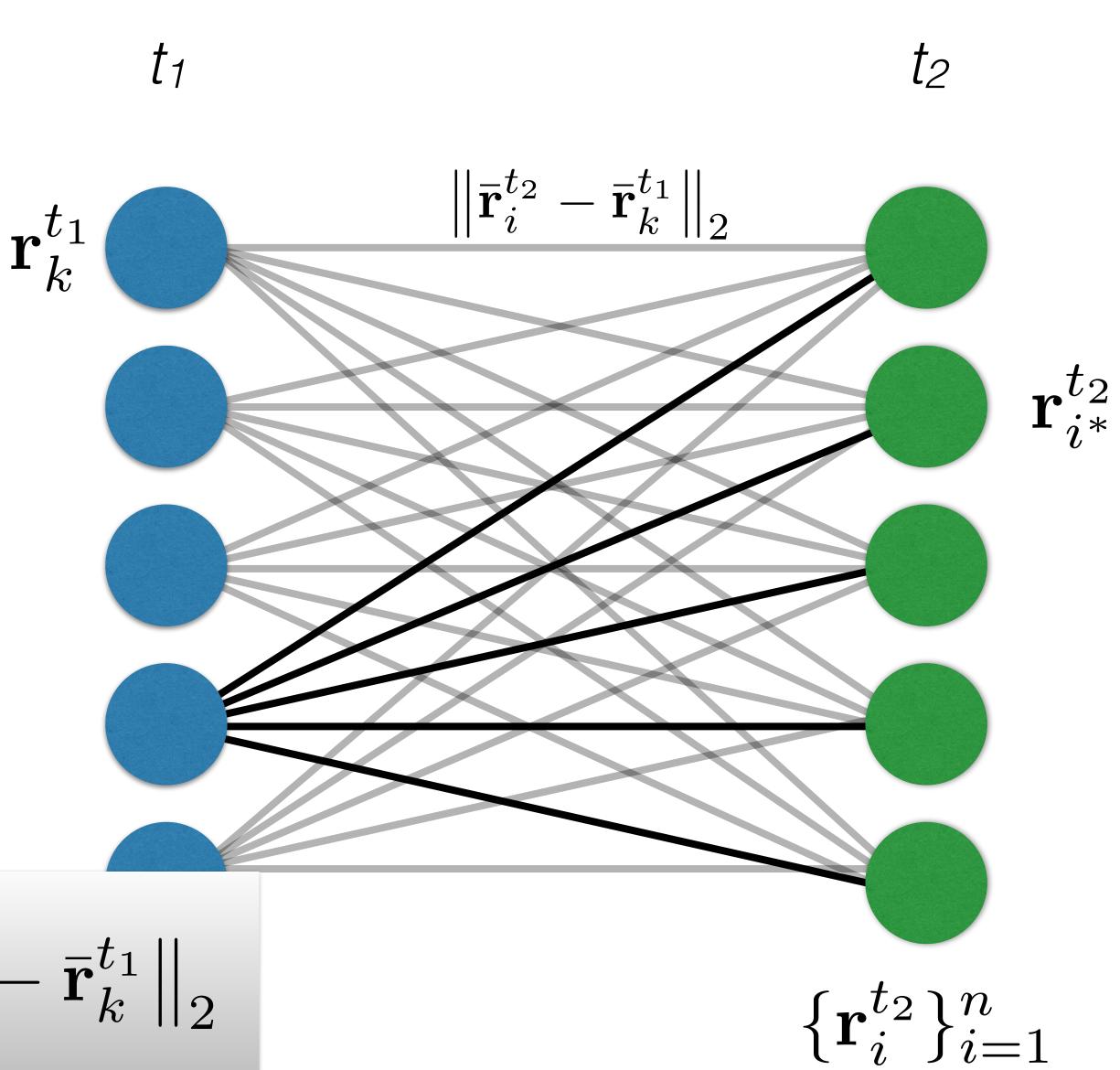
 $i^* = \underset{i}{\operatorname{arg\,min}} \|\bar{\mathbf{r}}_i^{t_2} - \bar{\mathbf{r}}_k^{t_1}\|_2$





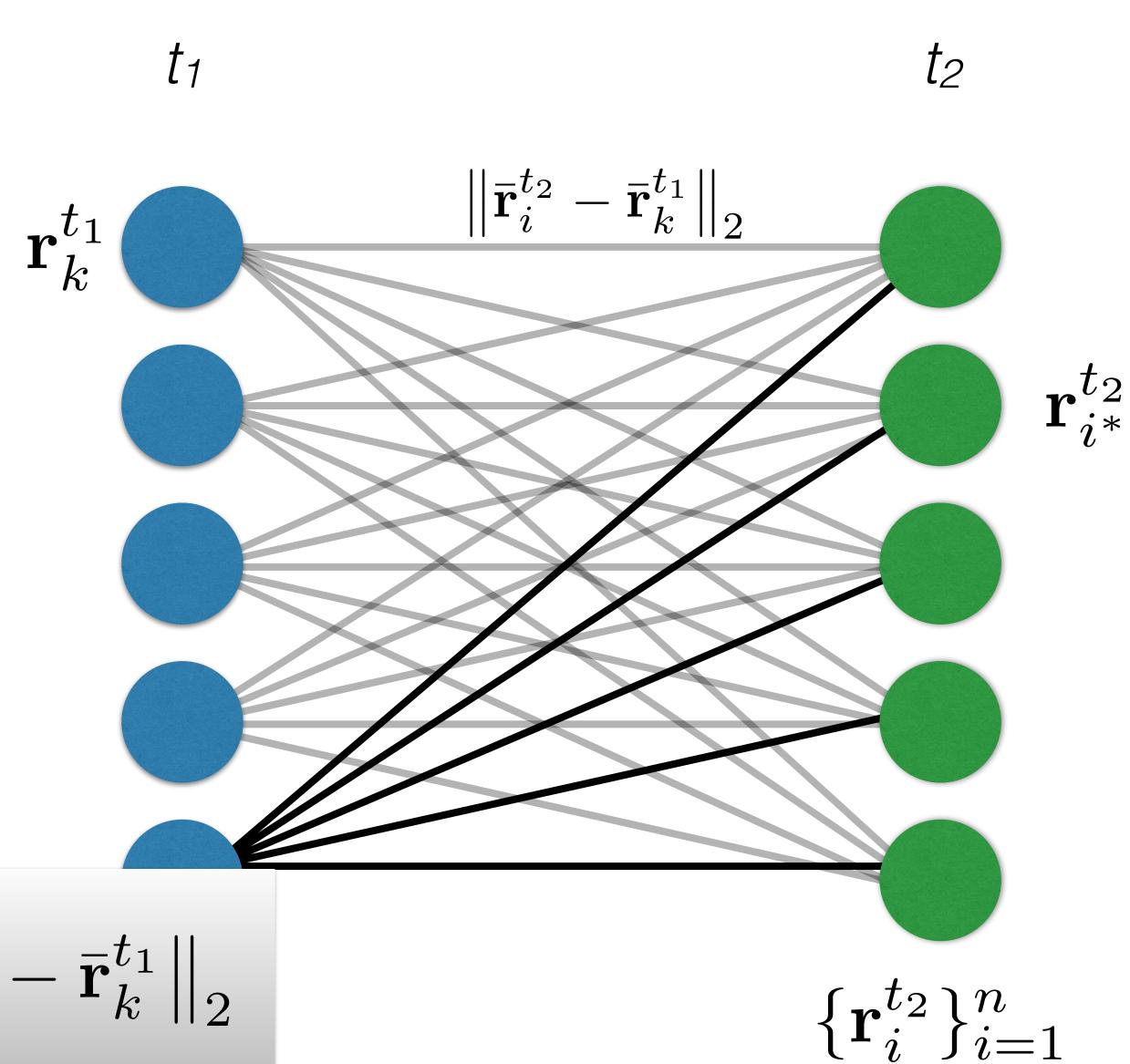
 $i^* = \underset{i}{\operatorname{arg\,min}} \|\bar{\mathbf{r}}_i^{t_2} - \bar{\mathbf{r}}_k^{t_1}\|_2$





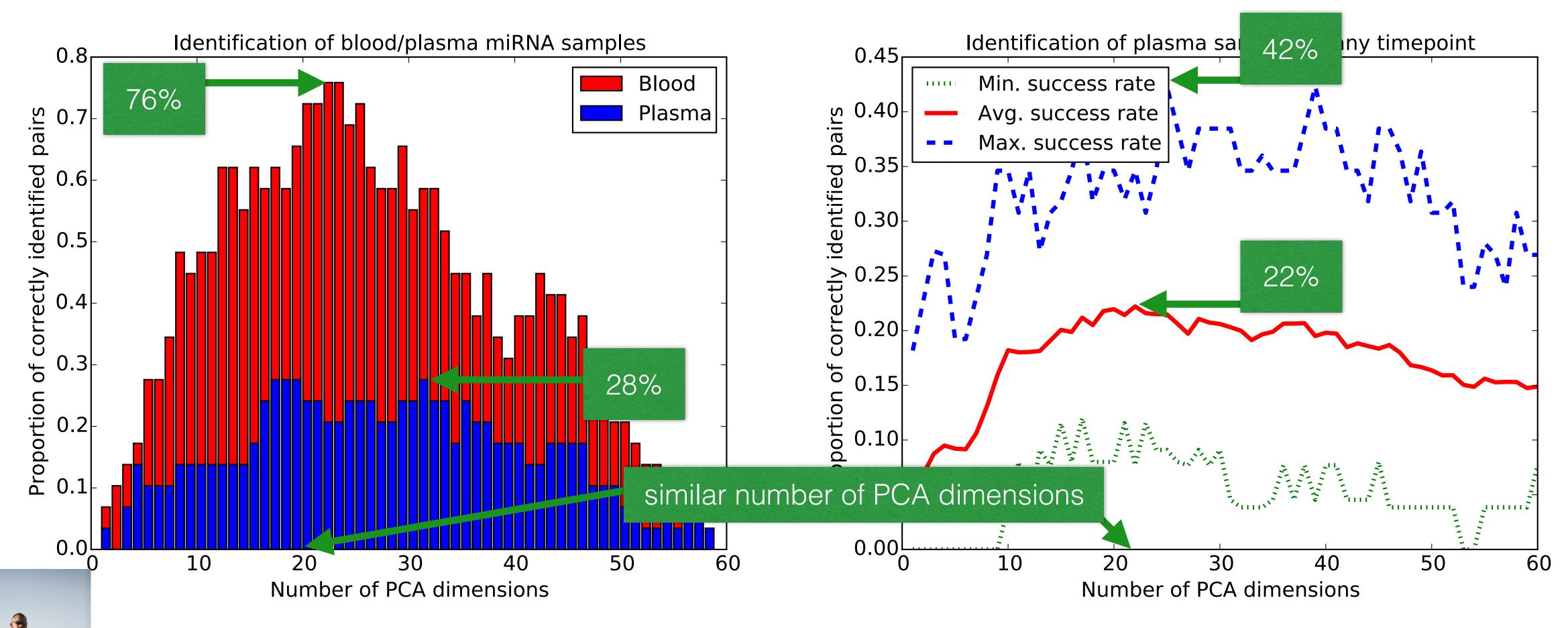
 $i^* = \underset{i}{\operatorname{arg\,min}} \|\bar{\mathbf{r}}_i^{t_2} - \bar{\mathbf{r}}_k^{t_1}\|_2$





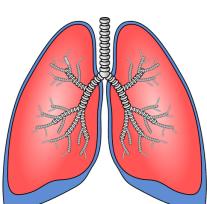
 $i^* = \underset{i}{\operatorname{arg\,min}} \|\bar{\mathbf{r}}_i^{t_2} - \bar{\mathbf{r}}_k^{t_1}\|_2$

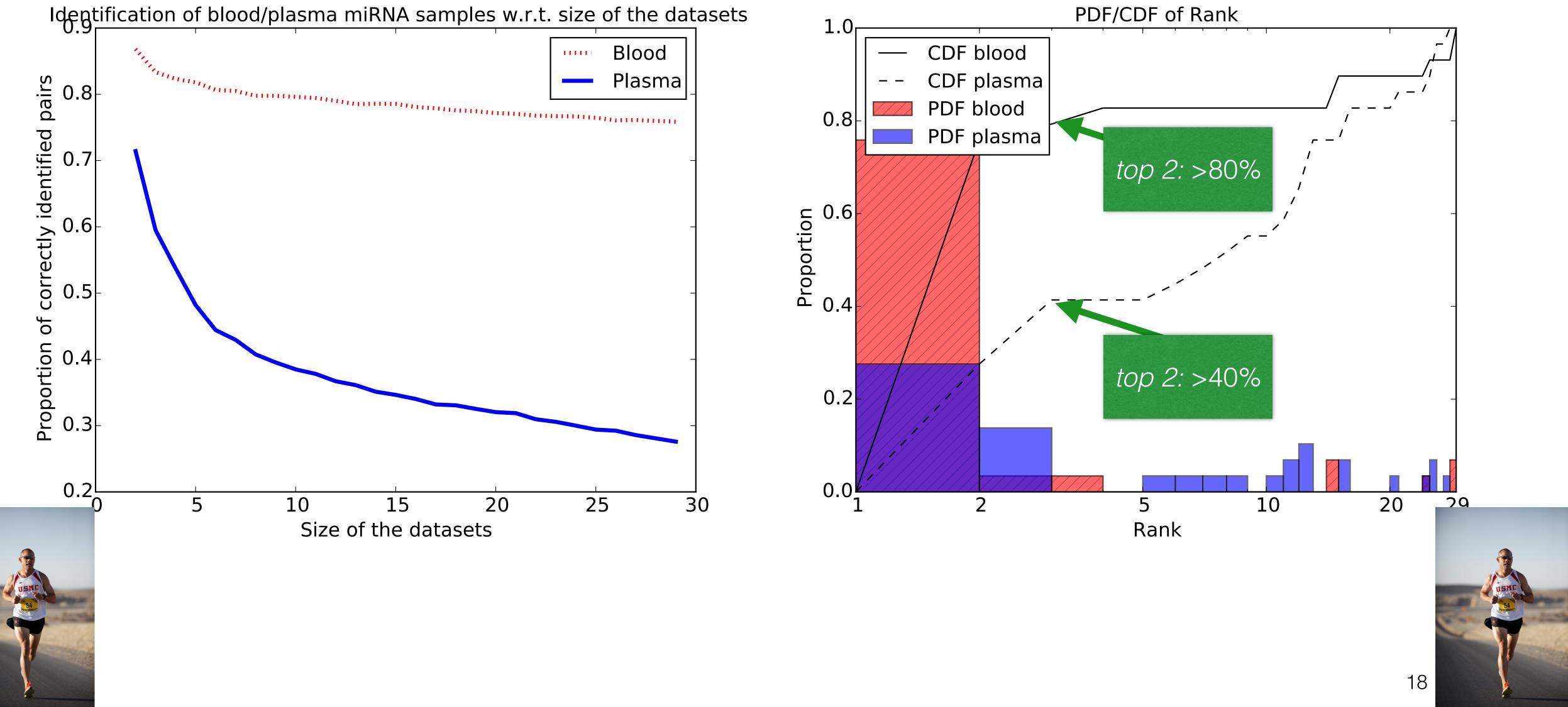




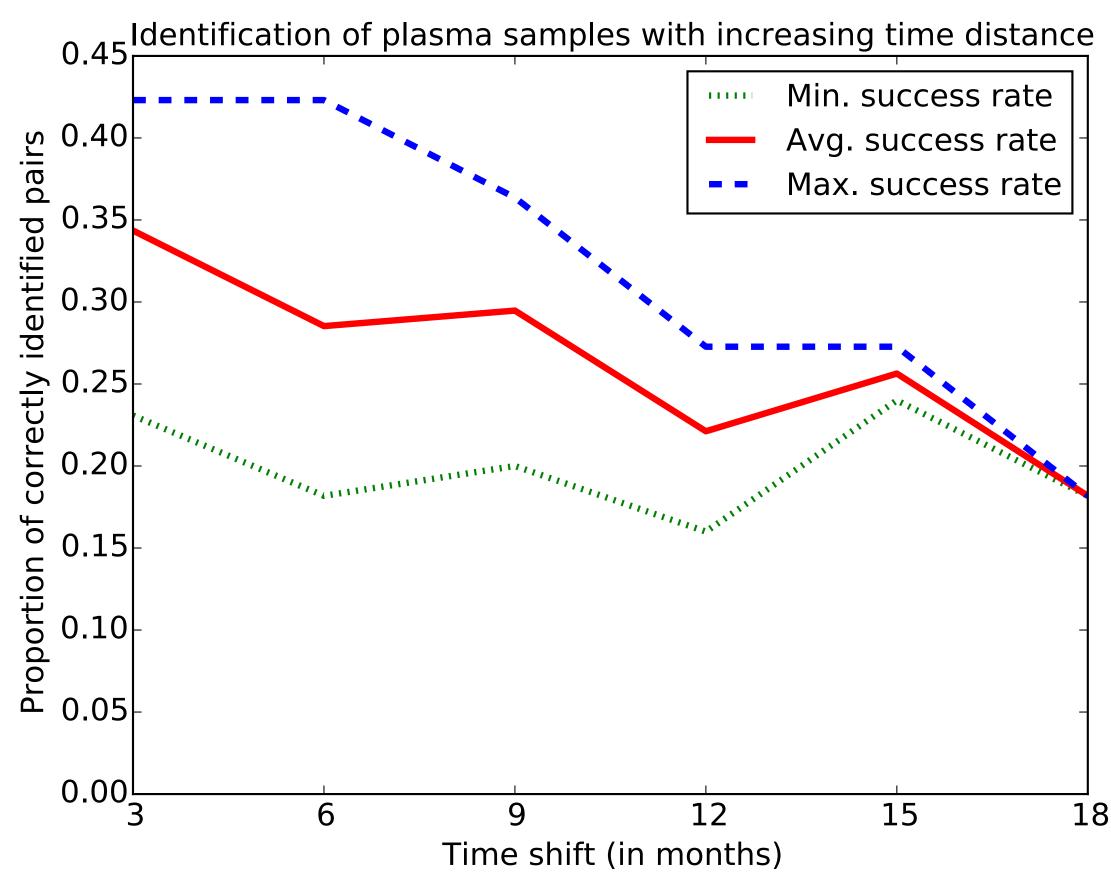
80% overlap in top10 miRNAs of first PCA component

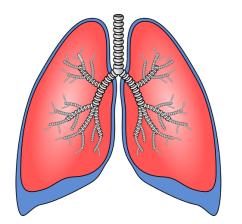




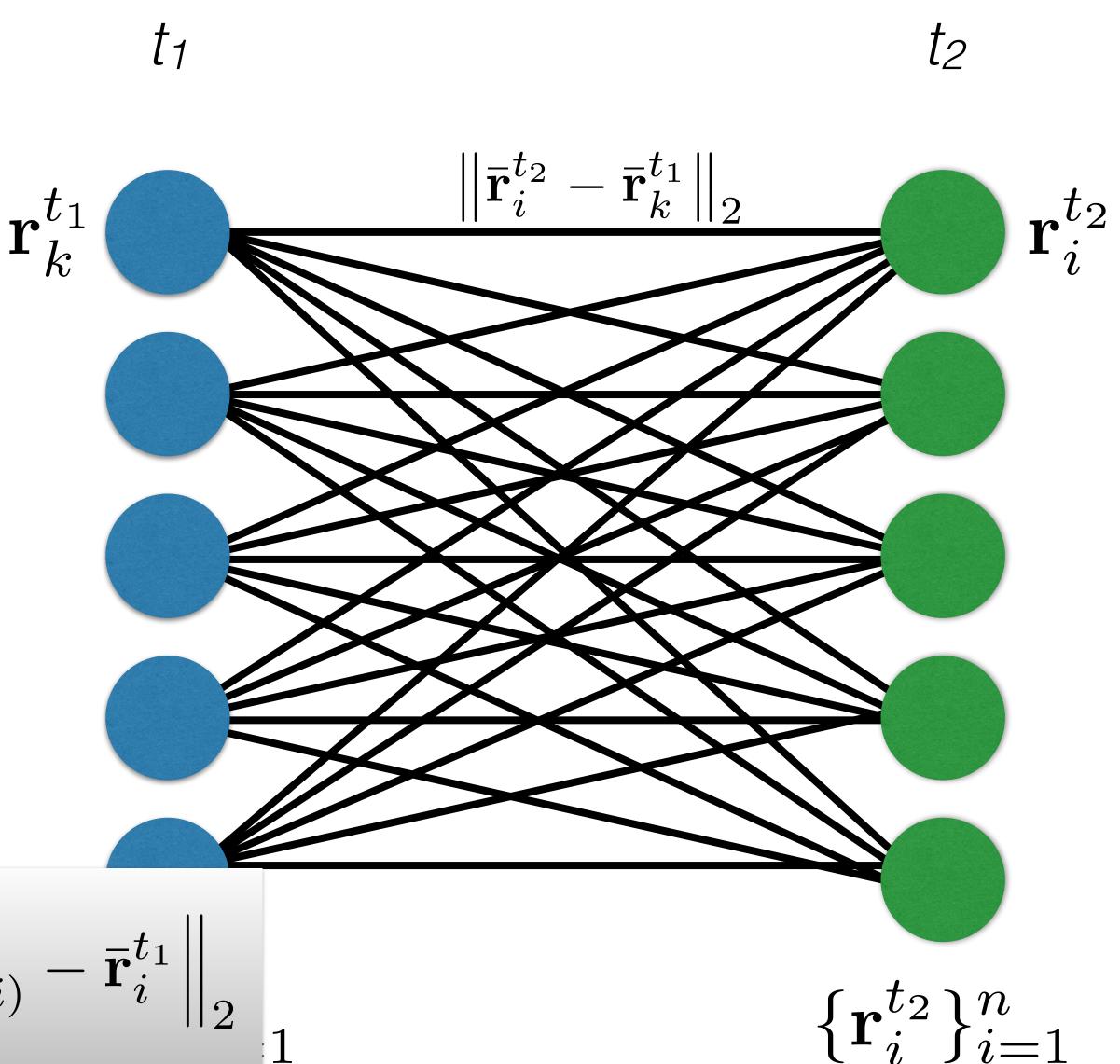








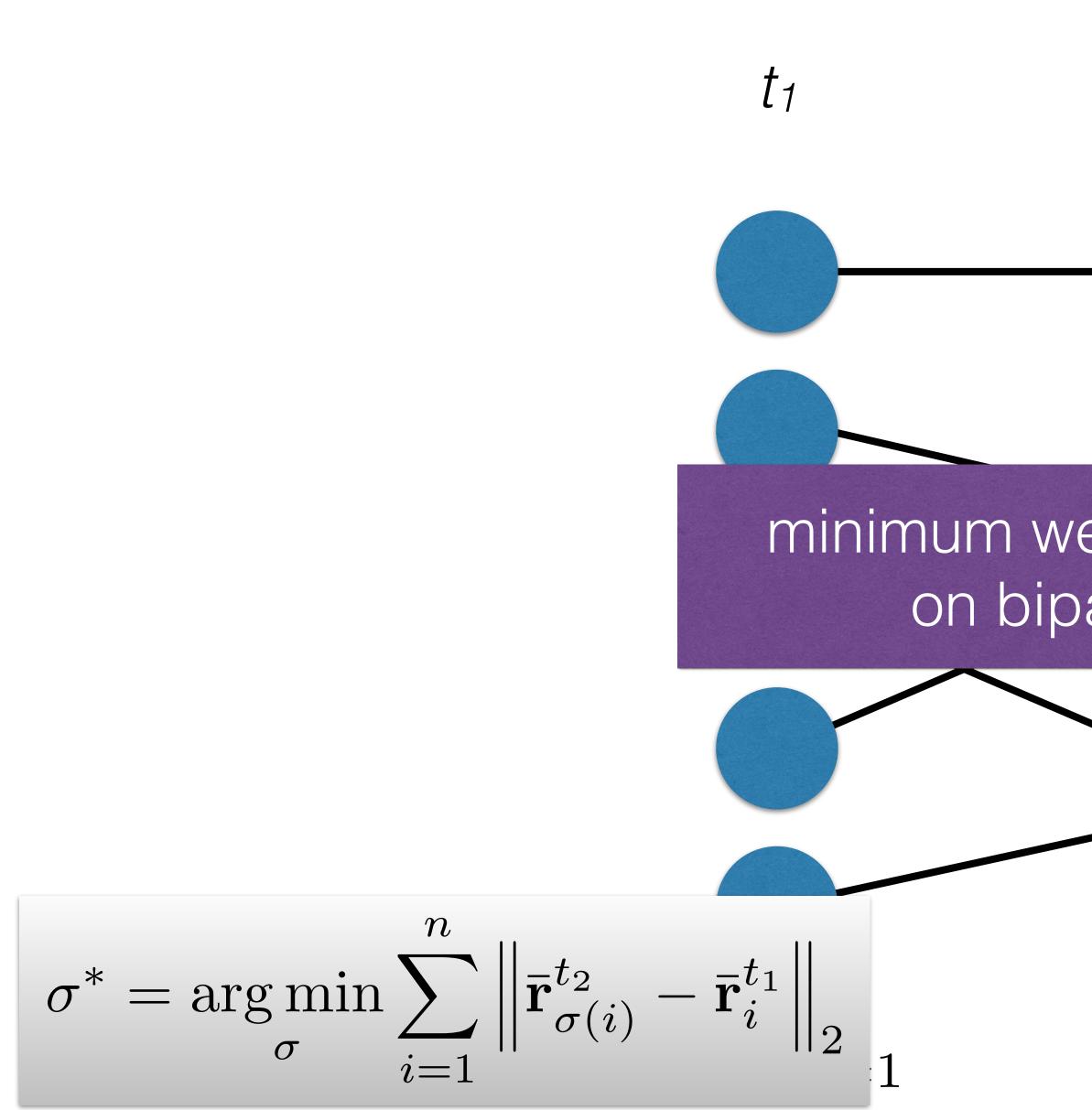




 $\sigma^* = \underset{\sigma}{\operatorname{arg\,min}} \sum_{i=1}^{n} \left\| \bar{\mathbf{r}}_{\sigma(i)}^{t_2} - \bar{\mathbf{r}}_{i}^{t_1} \right\|_2$



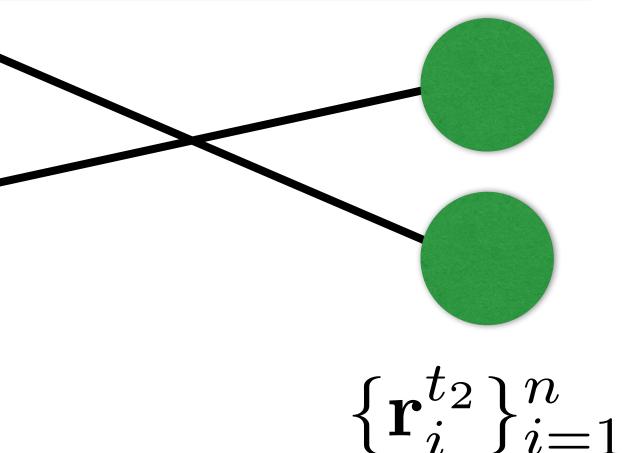


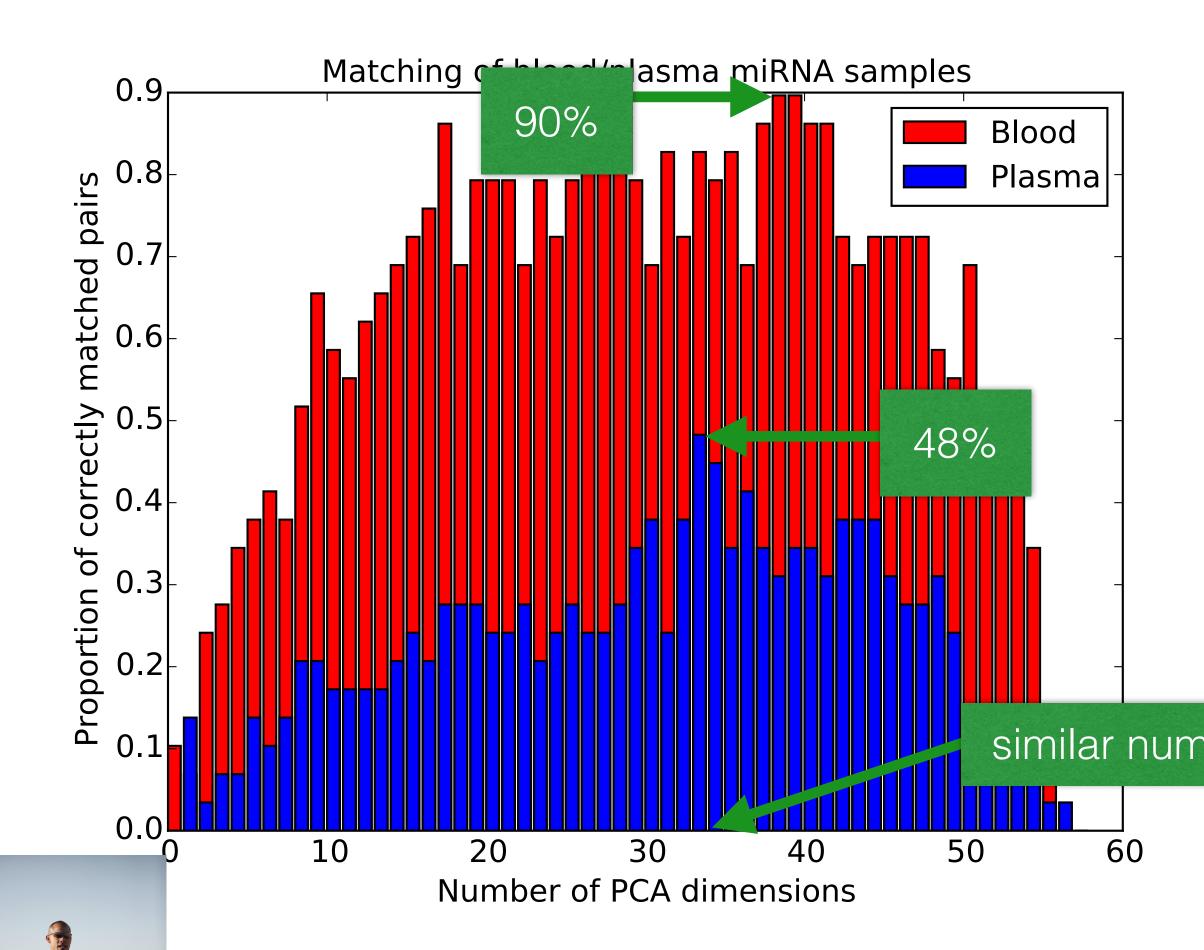




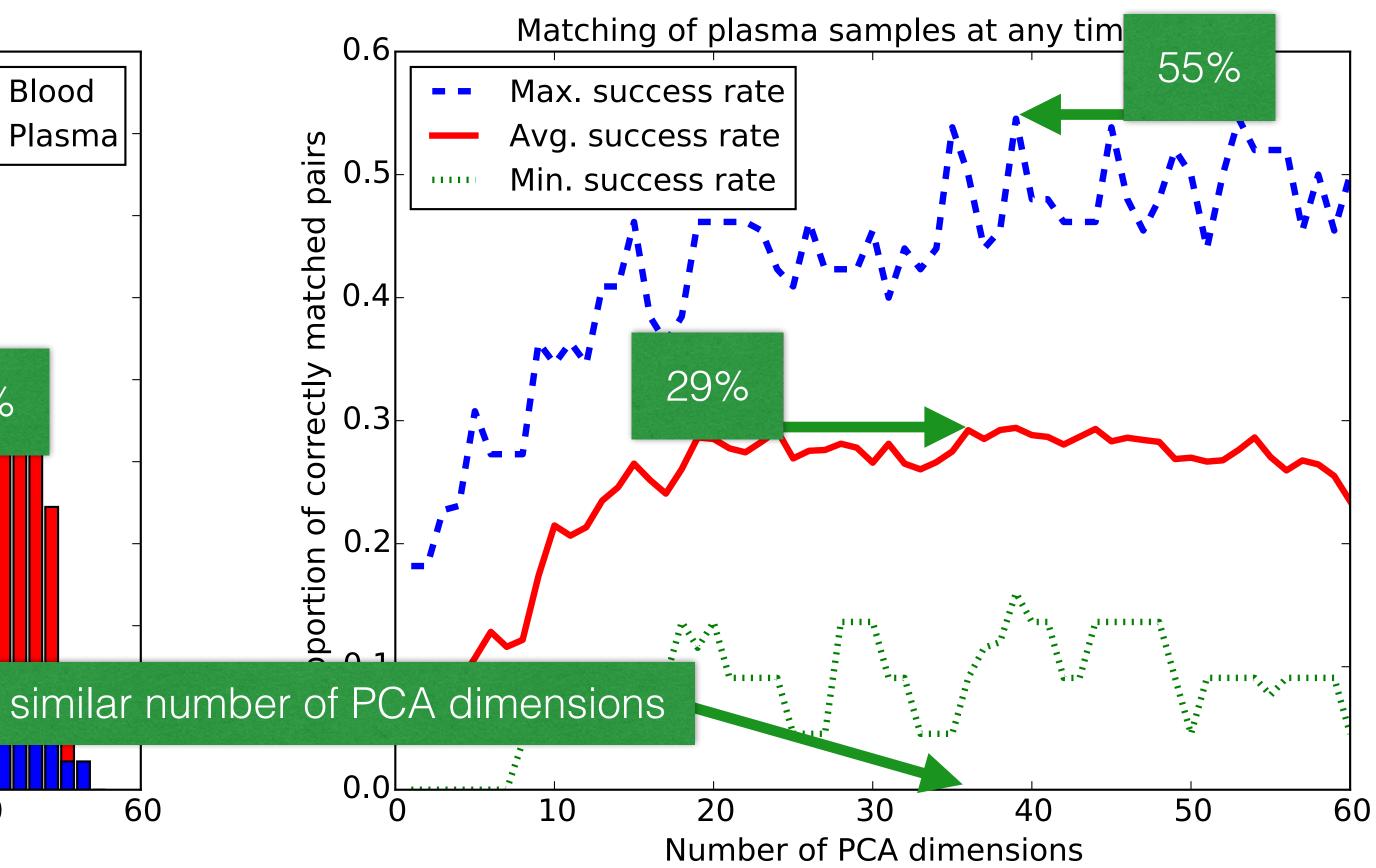
minimum weight assignment on bipartite graph

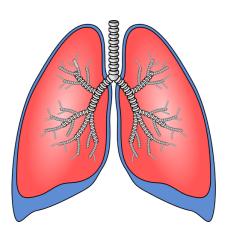
 t_2

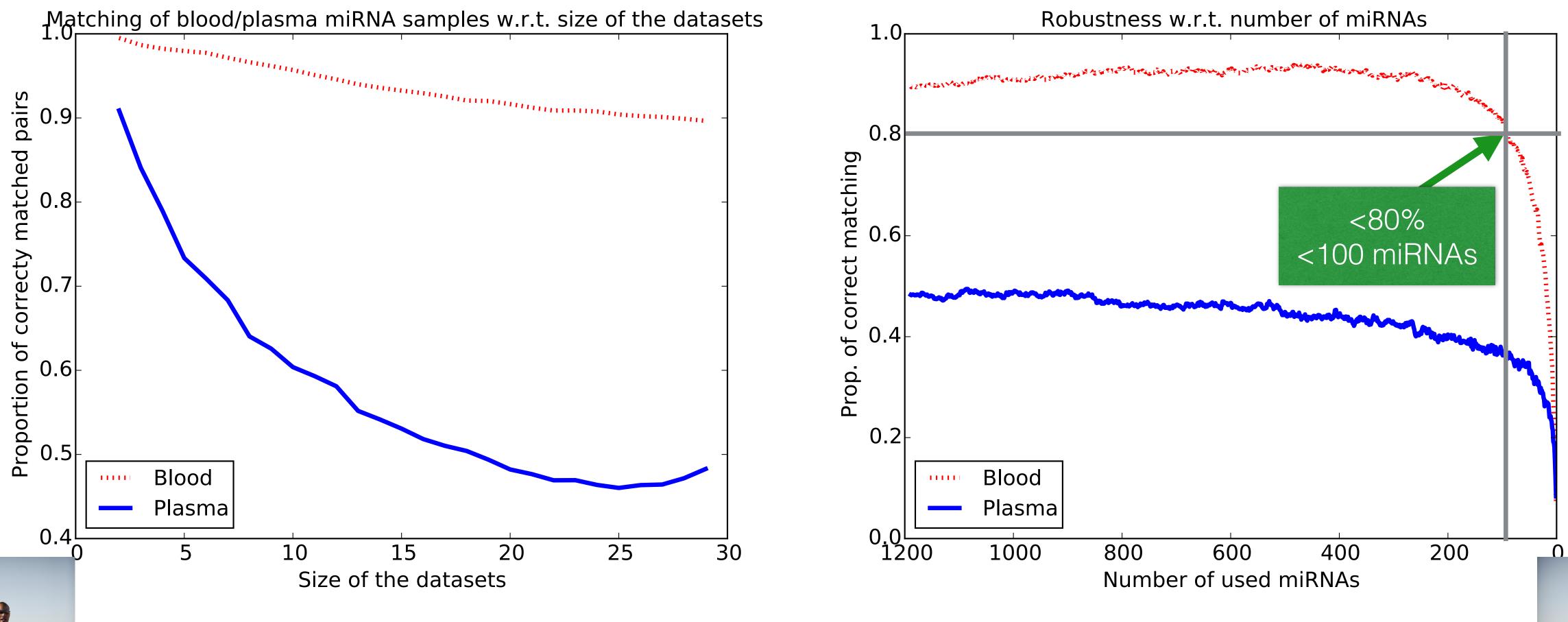








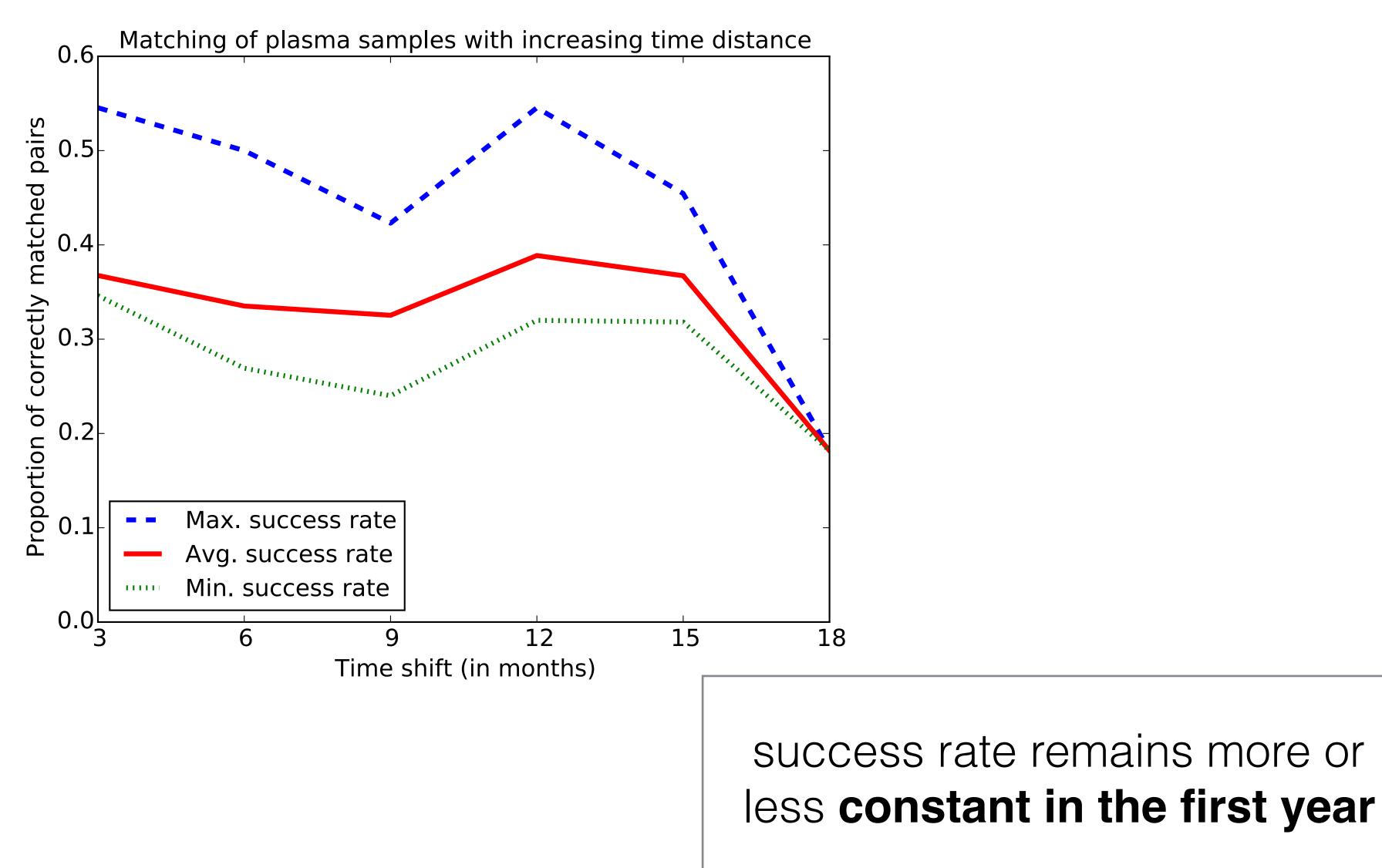


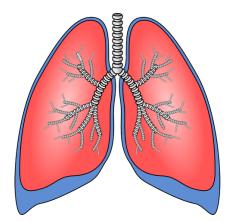








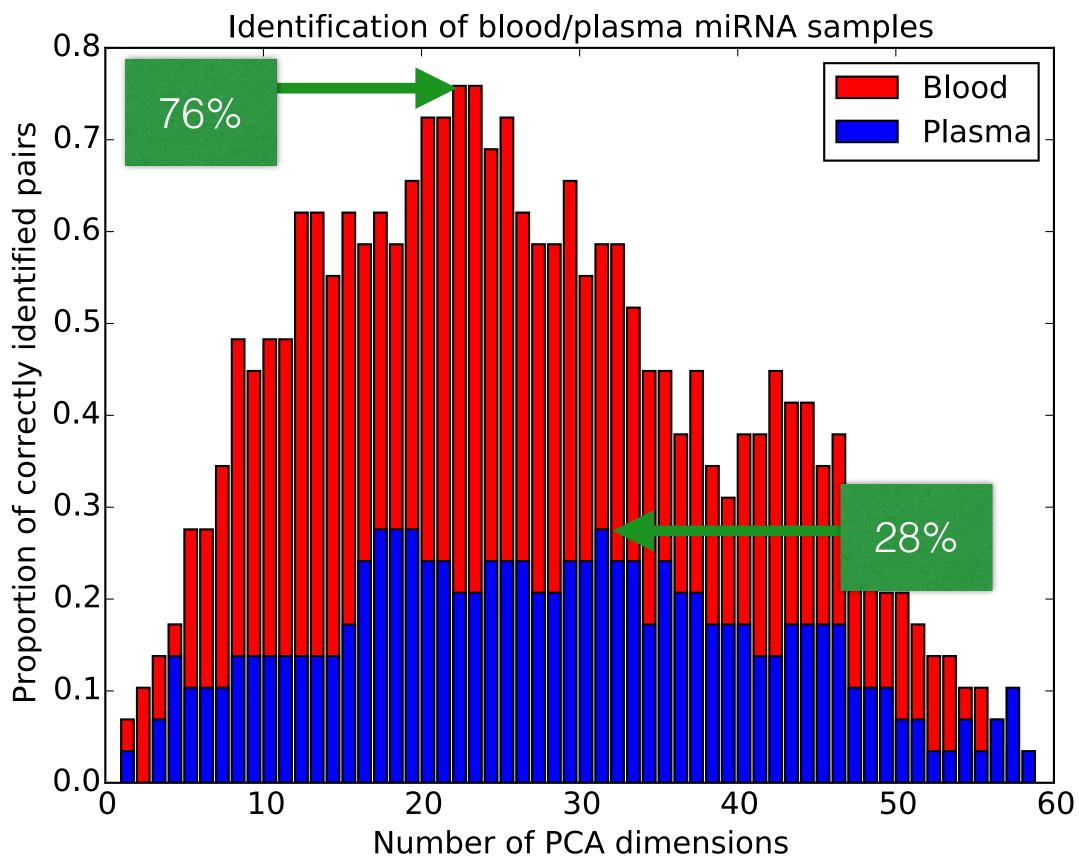






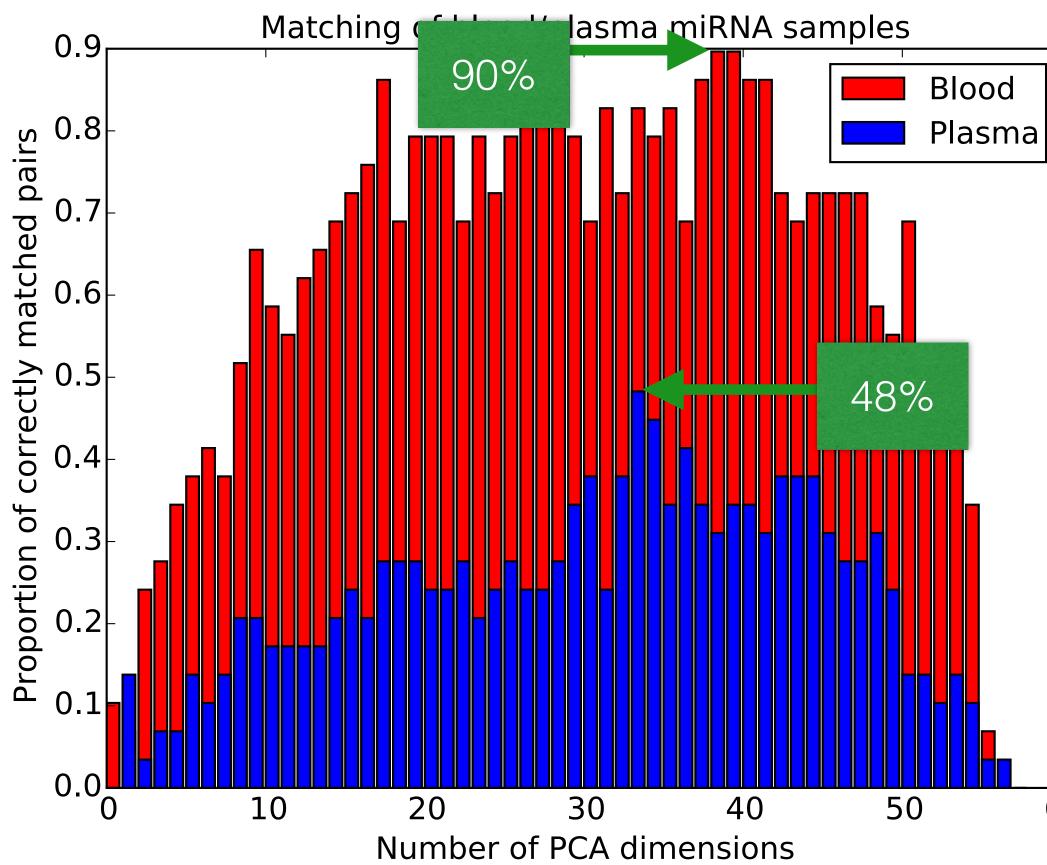








Matching Attack

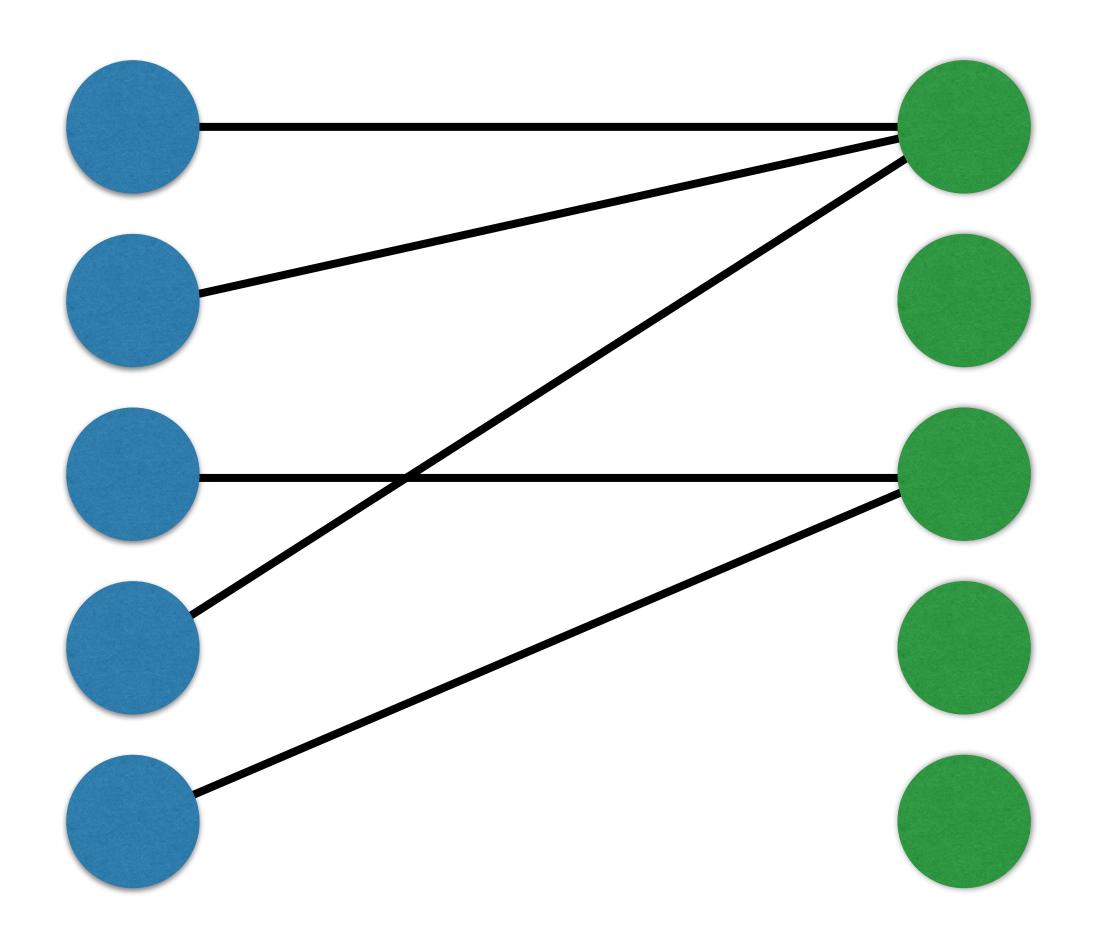






Downside of Identification Attack

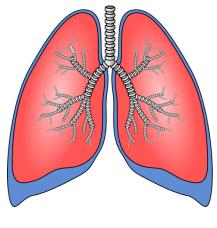
t1

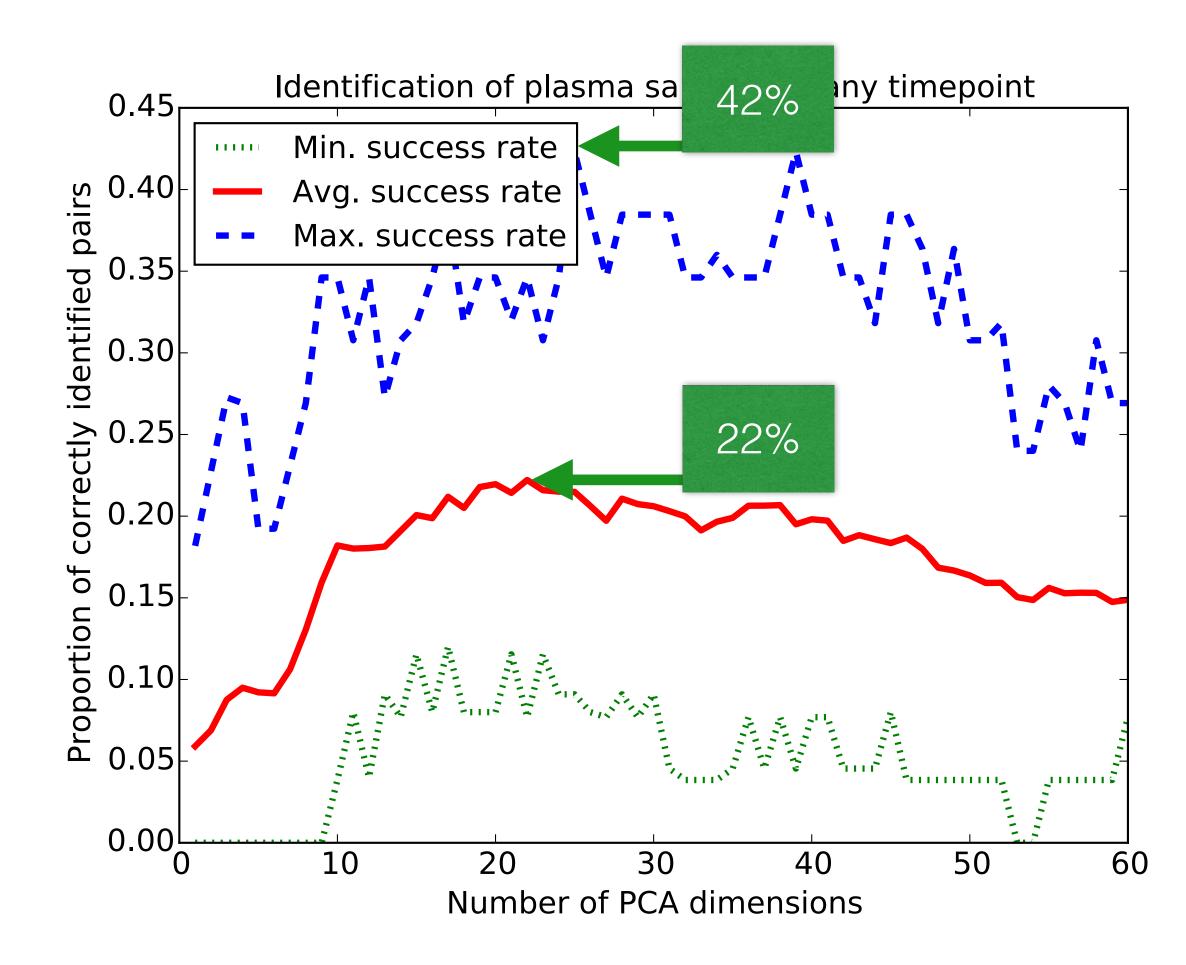




 t_2

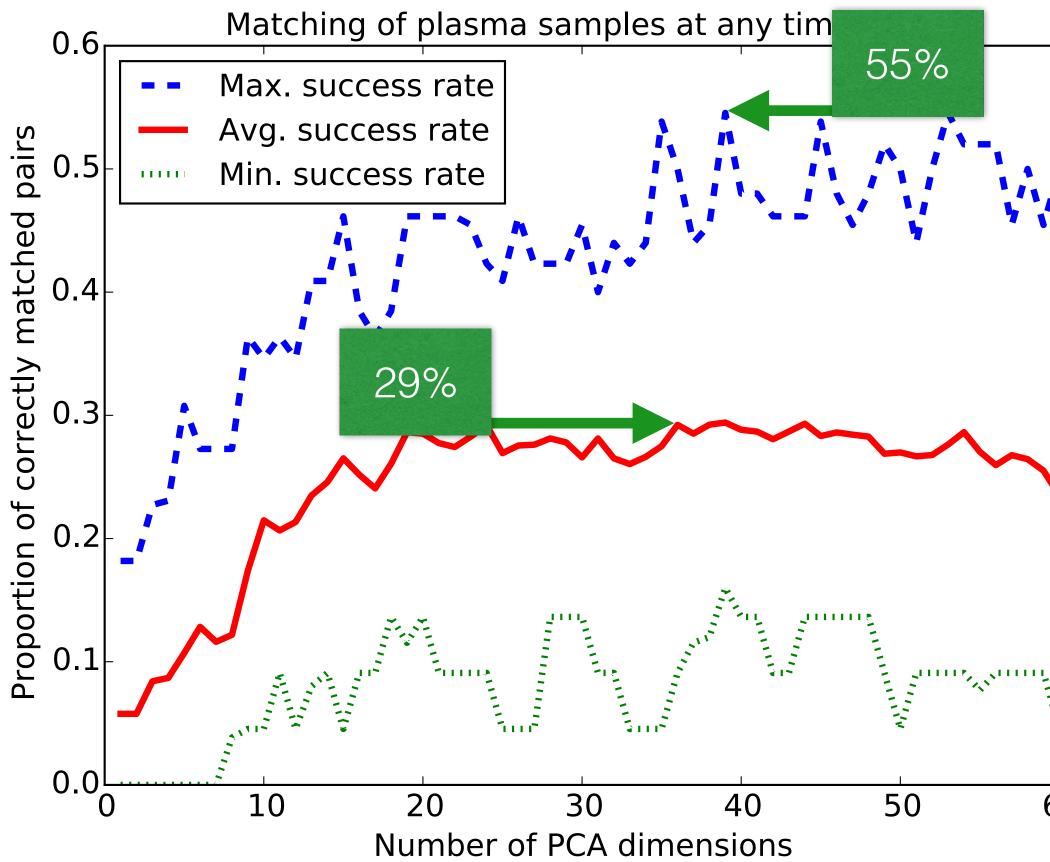








Matching Attack

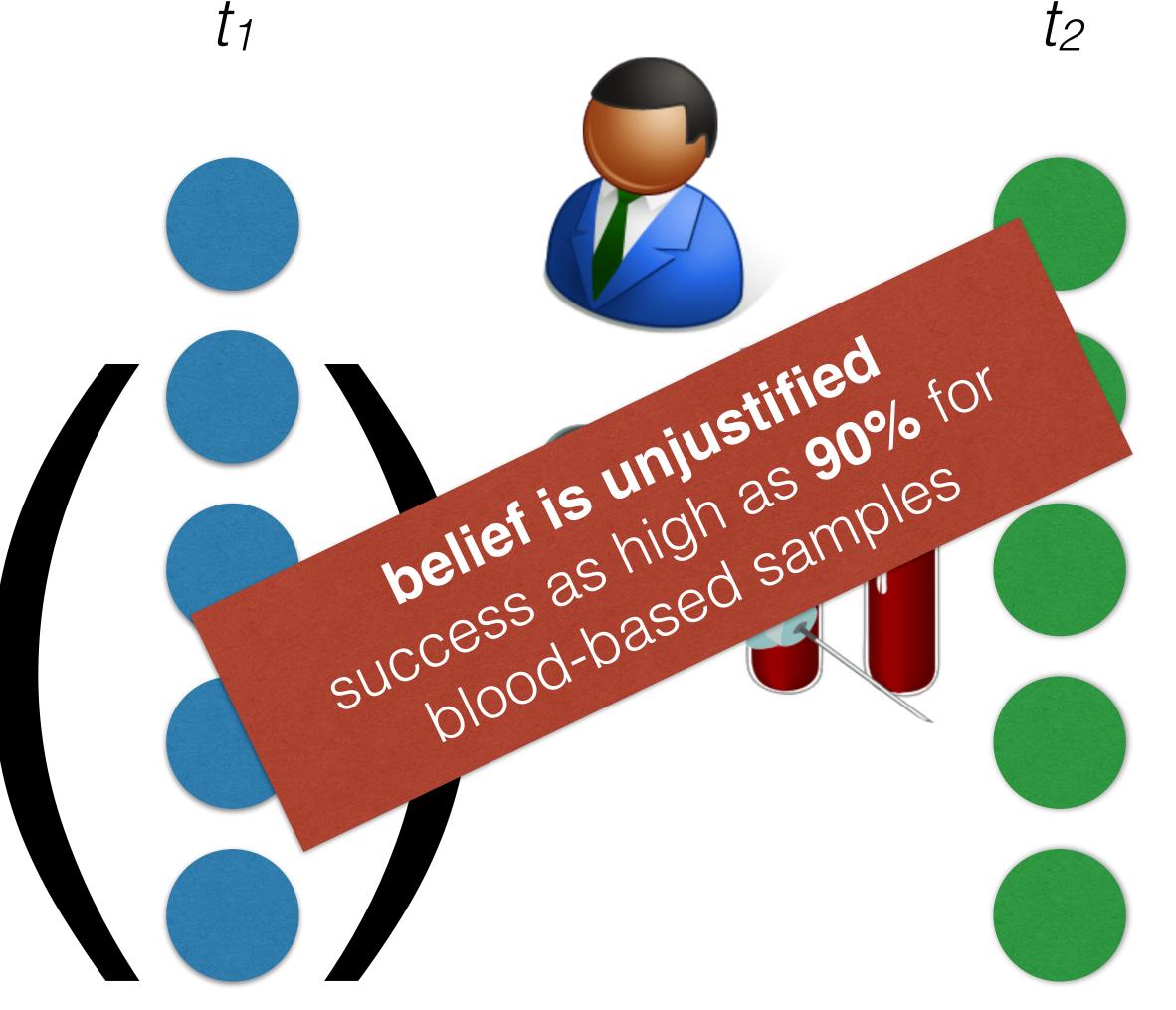






Common belief: no privacy threats from miRNAs, because of temporal variability

t₁







belief is unjustified linkability as high as **90%** for blood-based samples



there in fact are **privacy threats** inherent to epigenetic data

matching is more successful than identification





